

IN THE UNITED STATES
PATENT AND TRADEMARK OFFICE

In re Application of :

HIDENORI OHKI et al.

Serial No. : 09/248,267

Art Unit : 1654

Filed : February 11, 1999

Examiner : MARSHALL , S

For : CYCLIC HEXAPEPTIDES HAVING ANTIBIOTIC ACTIVITY

DECLARATION

I, FUMIAKI IKEDA, a citizen of Japan residing at No. 5-1-11, Kisaichi, Katano-shi, OSAKA, JAPAN, declare and say that :

I graduated from Osaka University, Faculty of Pharmacy, in March 1979;

I received the degree of Doctor of Pharmacy from Toho University, in 1987;

Since April 1979 , I have been, as a microbiologist, in the continuous employ of Fujisawa Pharmaceutical Co., Ltd., OSAKA, JAPAN and am now a Senior Research Manager of Medical Biology Research Laboratories;

I received from Mr. Hidenori Ohki the Test compounds 1 to 29, A and B, and I have conducted the anti-microbial tests.

Test (Anti-microbial activity) :

Test Method

In vitro antimicrobial activity of the test compounds was determined by the two-fold agar-plate dilution method.

One loopful of an overnight culture of each test microorganism in Sabourad broth containing 2 % Glucose (10^5 viable cells per ml) was streaked on Sabourad dextrose agar (SDA: dextrose 2%) containing graded concentrations of the test compounds, and the minimal inhibitory concentration (MIC) was expressed in term of $\mu\text{g/ml}$ after incubation at 30°C for 24 hours.

Test Organism : *C.albicans* FP633

Test Result

No.	MIC ($\mu\text{g/ml}$)
A	0.78
B	0.78
1	0.1
2	0.2
3	0.2
4	0.2
5	0.1
6	0.1
7	0.1
8	0.1
9	0.1
10	0.1
11	0.1
12	0.05
13	0.05
14	0.1
15	0.2
16	0.39
17	0.1
18	0.39

No.	MIC (µg/ml)
19	0.39
20	0.2
21	0.2
22	0.2
23	0.05
24	0.1
25	0.1
26	0.2
27	0.2
28	0.39
29	0.39

It is declared by the undersigned that all statements made herein of undersigned's own knowledge are true and that all statements made on information and belief are believed to be true, and further that these statements were made with the knowledge that willful false statements, and the like so made are punishable by fine or imprisonment, or both, under 18 U.S.C. 1001, and that such willful false statements may jeopardize the validity of this application or any patent issuing thereon.

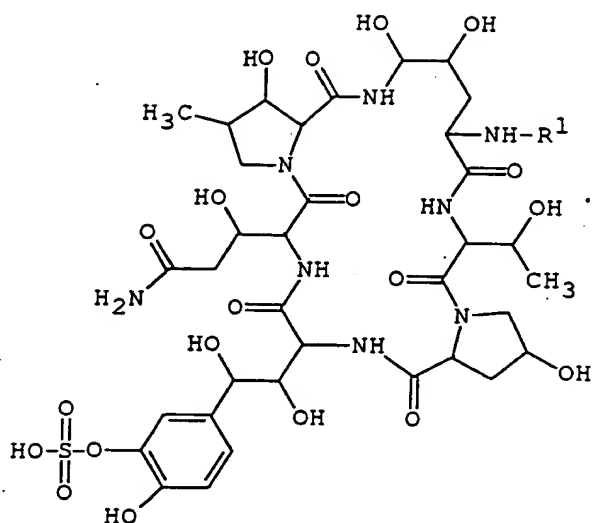
Fumiaki Ikeda
Fumiaki Ikeda

This 1st day of April, 1999
OSAKA, JAPAN

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CLAIMS

1. A polypeptide compound of the following general formula :



wherein R¹ is lower alkanoyl substituted with

Ex 44 { unsaturated 6-membered heteromonocyclic group containing at least one nitrogen atom which may have one or more suitable substituent(s);

~~lower alkanoyl substituted with 1,2,3,4-tetrahydroisoquinoline which may have one or more suitable substituent(s);~~

Ex 45 { lower alkanoyl substituted with unsaturated condensed heterocyclic group containing at least one oxygen atom which may have one or more suitable substituent(s);

Ex 12 { lower alkanoyl substituted with unsaturated condensed heterocyclic group containing 1 to 3 sulfur atom(s)

which may have one or more suitable
substituent(s);

~~lower alkanoyl substituted with
unsaturated condensed heterocyclic
group containing 2 or more nitrogen
atom(s) which may have one or more
suitable substituent(s);~~

EX 26

lower alkanoyl substituted with
saturated 3 to 8 membered
heteromonocyclic group containing at
least one nitrogen atom which may have
one or more suitable substituent(s);

EX 84

ar(lower)alkenoyl substituted with
aryl which may have one or more
suitable substituent(s);

EX 21

naphthyl(lower)alkenoyl which may
have one or more higher alkoxy;

EX 31

lower alkynoyl which may have one or
more suitable substituent(s);

EX 20

(C₂-C₆)alkanoyl substituted with
naphthyl having higher alkoxy;

EX 16

ar(C₂-C₆)alkanoyl substituted with
aryl having one or more suitable
substituent(s), in which ar(C₂-C₆)-
alkanoyl may have one or more suitable
substituent(s);

EX 40

aroyl substituted with heterocyclic
group which may have one or more
suitable substituent(s), in which aroyl
may have one or more suitable
substituent(s);

~~aroyl substituted with aryl having
heterocyclic(higher)alkoxy, in which
heterocyclic group may have one or more
suitable substituent(s);~~

EX24 (aroyl substituted with aryl having
lower alkoxy(higher)alkoxy;

~~aroyl substituted with aryl having
lower alkenyl(lower)alkoxy;
aroyl substituted with 2 lower
alkoxy;~~

EX22 (aroyl substituted with aryl having
lower alkyl;

EX35 (aroyl substituted with aryl having
higher alkyl;

~~aryloxy(lower)alkanoyl which may have
one or more suitable substituent(s);~~

EX8 (ar(lower)alkoxy(lower)alkanoyl which
may have one or more suitable
substituent(s);

~~aryl amino(lower)alkanoyl which may
have one or more suitable
substituent(s);~~

EX54 (lower alkanoyl substituted with
pyrazolyl which has lower alkyl and
aryl having higher alkoxy;

EX118 (lower alkoxy(higher)alkanoyl, in
which higher alkanoyl may have one or
more suitable substituent(s);

~~aroyl substituted with aryl having
heterocyclicoxy, in which
heterocyclicoxy may have one or more
suitable substituent(s);~~

EX74 (aroyl substituted with
cyclo(lower)alkyl having lower alkyl;
EX113 indolylcarbonyl having higher alkyl;
EX82 naphthoyl having lower alkyl;
EX81 naphthoyl having higher alkyl;

~~naphthoyl having lower
alkoxy(higher)alkoxy;~~

EX103 (aroyl substituted with aryl having
lower alkoxy(lower)alkoxy(higher)-
alkoxy;

EX114 (aroyl substituted with aryl having
lower alkoxy(lower)alkoxy;

~~aroyl substituted with aryl which has
aryl having lower alkoxy;~~

EX89 (aroyl substituted with aryl which has
aryl having lower alkoxy(lower)alkoxy;

EX112 (aroyl substituted with aryl having
heterocyclicoxy(higher)alkoxy;

EX51 (aroyl substituted with aryl having
aryloxy(lower)alkoxy;

~~aroyl substituted with aryl having
heterocycliccarbonyl(higher)alkoxy;~~

EX109 (lower alkanoyl substituted with
oxazolyl which has aryl having higher
alkoxy;

~~lower alkanoyl substituted with furyl
which has aryl substituted with aryl
having lower alkoxy;
lower alkanoyl substituted with
triazolyl which has oxo and aryl having
higher alkyl;~~

EX62 higher alkanoyl having hydroxy;

EX60 (higher alkanoyl having ar(lower)alkyl
and hydroxy; or

EX63 3-methyl-tridecenoyl; or

~~(C₂-C₆)alkanoyl substituted with aryl
having higher alkoxy, in which (C₂-C₆)-
alkanoyl may have amino or protected
amino, and~~

a pharmaceutically acceptable salt thereof.

2. A compound of claim 1, wherein